

SEQUENCE LISTING

<110> Altmann, Scott W
Wang, Luquan
Graziano, Michael
Murgolo, Nick

<120> NPC1L1 (NPC3) AND METHODS OF USE THEREOF

<130> JB01603K2 US

<140> 10/663,208

<141> 2003-09-16

<150> 60/397,442

<151> 2002-07-19

<150> 10/621,758

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					gcc Ala											2784
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	cta Leu 1280					gca Ala 1285										3879
	tac Tyr 1295					gat Asp 1300							tac Tyr	_		3924
	gcc Ala 1310	cca Pro	gaa Glu	ctt Leu	gcc Ala	cac His 1315	gga Gly	gct Ala	aat Asn	gct Ala	gct Ala 1320	aga Arg	agc Ser	tct Ser	,	3969
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Arg Pro Ser Phe Tyr Met Gly Arg Met Pro Gly Trp Leu Ala Leu Ile Ile Ile Phe Thr Ala Val Phe Val Leu Ser Ala Val Leu Val Arg Leu Arg Val Val Ser Asn Arg Asn Lys Asn Lys Ala Glu Gly Pro Gln Glu Ala Pro Lys Leu Pro His Lys His Lys Leu Ser Pro His Thr Ile Leu Gly Arg Phe Phe Gln Asn Trp Gly Thr Arg Val Ala Ser Trp Pro Leu Thr Val Leu Ala Leu Ser Phe Ile Val Val Ile Ala Leu Ala Ala Gly Leu Thr Phe Ile Glu Leu Thr Thr Asp Pro Val Glu Leu Trp Ser Ala Pro Lys Ser Gln Ala Arg Lys Glu Lys Ser Phe His Asp Glu His Phe Gly Pro Phe Phe Arg Thr Asn Gln Ile Phe Val Thr Ala Arg Asn Arg Ser Ser Tyr Lys Tyr Asp Ser Leu Leu Cly Ser Lys Asn Phe Ser Gly Ile Leu Ser Leu Asp Phe Leu Leu Glu Leu Glu Leu Gln Glu Arg Leu Arg His Leu Gln Val Trp Ser Pro Glu Ala Glu Arg Asn Ile Ser Leu Gln Asp Ile Cys Tyr Ala Pro Leu Asn Pro Tyr Asn Thr Ser Leu Ser Asp Cys Cys Val Asn Ser Leu Leu Gln Tyr Phe Gln Asn Asn Arg Thr Leu Leu Met Leu Thr Ala Asn Gln Thr Leu Asn Gly Gln

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- Arg Gln Glu Ala Ser Arg Pro Asp Val Leu Cys Cys Phe Ser Thr Arg
- Lys Leu Pro Pro Pro Lys Glu Lys Glu Gly Leu Leu Arg Phe Phe
- Arg Lys Ile Tyr Ala Pro Phe Leu Leu His Arg Phe Ile Arg Pro Val
- Val Met Leu Leu Phe Leu Thr Leu Phe Gly Ala Asn Leu Tyr Leu Met
- Cys Asn Ile Asn Val Gly Leu Asp Gln Glu Leu Ala Leu Pro Lys Asp
- Ser Tyr Leu Ile Asp Tyr Phe Leu Phe Leu Asn Arg Tyr Leu Glu Val
- Gly Pro Pro Val Tyr Phe Val Thr Thr Ser Gly Phe Asn Phe Ser Ser
- Glu Ala Gly Met Asn Ala Thr Cys Ser Ser Ala Gly Cys Lys Ser Phe
- Ser Leu Thr Gln Lys Ile Gln Tyr Ala Ser Glu Phe Pro Asp Gln Ser
- Tyr Val Ala Ile Ala Ala Ser Ser Trp Val Asp Asp Phe Ile Asp Trp
- Leu Thr Pro Ser Ser Cys Cys Arg Leu Tyr Ile Arg Gly Pro His
- Lys Asp Glu Phe Cys Pro Ser Thr Asp Thr Ser Phe Asn Cys Leu Lys
- Asn Cys Met Asn Arg Thr Leu Gly Pro Val Arg Pro Thr Ala Glu Gln

Phe His Lys Tyr Leu Pro Trp Phe Leu Asn Asp Pro Pro Asn Ile Arg Cys Pro Lys Gly Gly Leu Ala Ala Tyr Arg Thr Ser Val Asn Leu Ser Ser Asp Gly Gln Val Ile Ala Ser Gln Phe Met Ala Tyr His Lys Pro Leu Arg Asn Ser Gln Asp Phe Thr Glu Ala Leu Arg Ala Ser Arg Leu Leu Ala Ala Asn Ile Thr Ala Asp Leu Arg Lys Val Pro Gly Thr Asp Pro Asn Phe Glu Val Phe Pro Tyr Thr Ile Ser Asn Val Phe Tyr Gln Gln Tyr Leu Thr Val Leu Pro Glu Gly Ile Phe Thr Leu Ala Leu Cys Phe Val Pro Thr Phe Val Val Cys Tyr Leu Leu Gly Leu Asp Met Cys Ser Gly Ile Leu Asn Leu Leu Ser Ile Ile Met Ile Leu Val Asp Thr Ile Gly Leu Met Ala Val Trp Gly Ile Ser Tyr Asn Ala Val Ser Leu Ile Asn Leu Val Thr Ala Val Gly Met Ser Val Glu Phe Val Ser His Ile Thr Arq Ser Phe Ala Val Ser Thr Lys Pro Thr Arg Leu Glu Arg Ala Lys Asp Ala Thr Val Phe Met Gly Ser Ala Val Phe Ala Gly Val Ala Met Thr Asn Phe Pro Gly Ile Leu Ile Leu Gly Phe Ala Gln Ala

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		_	_		_	_	cgc Arg				_		_	624
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							att Ile							720
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							gcc Ala 265							816
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								atc Ile								1104
								acg Thr								1152
								gag Glu		_			_	_		1200
								cag Gln								1248
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								gcc Ala								1440
								agc Ser								1488
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	aag Lys	_				_		_	_		_	_				1728
	aat Asn			_		_		_	_	_	_	_	_	_		1776
	gag Glu	_				_	_	_	_		_	_		_	_	1824
	atg Met 610		_	_	_		_	_		_		_	_	_		1872
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	agc Ser															2016
	gcc Ala															2064
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													gag Glu			2832
													att Ile			2880
													ccc Pro			2928
													cta Leu 990			2976
			atc Ile					Va]					l G	_	ag ttc In Phe	3024
	aag Lys 101	Ty:	t cti r Lei				e Le				g Pı		aac a Asn I	-		3069

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						tac Tyr 1105							G1y 999		3339
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Leu Met Thr Leu Ser Asn Val Ser Cys Leu Ser Asn Thr Pro Ala Arg 50 55 60

Lys Ile Thr Gly Asp His Leu Ile Leu Leu Gln Lys Ile Cys Pro Arg 70 75 80

Leu Tyr Thr Gly Pro Asn Thr Gln Ala Cys Cys Ser Ala Lys Gln Leu 85 90 95

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Val	Pro	Ala	Ala 180	Ala	Thr	Leu	Ala	Val 185	Gly	Thr	Met	Cys	Gly 190	Val	Tyr
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Thr	Gly 210	Asn	Gly	Leu	Ala	Pro 215	Leu	Asp	Ile	Thr	Phe 220	His	Leu	Leu	Glu
Pro 225	Gly	Gln	Ala	Val	Gly 230	Ser	Gly	Ile	Gln	Pro 235	Leu	Asn	Glu	Gly	Val 240
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Phe Gly P	Pro Phe Ph 40		Asn Gl	n Val Ile 410	Leu Thr	Ala Pro 415	
Arg Ser S	Ser Tyr Ar 420	g Tyr Asp	Ser Le 42		Gly Pro	Lys Asn 430	Phe
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Glu Arg L 450	eu Arg Hi	s Leu Gln 455	-	p Ser Pro	Glu Ala 460	Gln Arg	Asn
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Pro Leu T 530	Thr Phe Ly	s Asp Gly 535		a Leu Ala	Leu Ser 540	Cys Met	Ala
Asp Tyr G 545	Sly Ala Pr	o Val Phe 550	Pro Ph	e Leu Ala 555	-	Gly Tyr	Lys 560
Gly Lys A	Asp Tyr Se 56		Glu Al	a Leu Ile 570	Met Thr	Phe Ser	
Asn Asn T	Tyr Pro Al 580	a Gly Asp	Pro Ar	-	Gln Ala	Lys Leu 590	Trp

Glu Glu Ala Phe Leu Glu Glu Met Arg Ala Phe Gln Arg Arg Met Ala Gly Met Phe Gln Val Thr Phe Thr Ala Glu Arg Ser Leu Glu Asp Glu Ile Asn Arg Thr Thr Ala Glu Asp Leu Pro Ile Phe Ala Thr Ser Tyr Ile Val Ile Phe Leu Tyr Ile Ser Leu Ala Leu Gly Ser Tyr Ser Ser Trp Ser Arg Val Met Val Asp Ser Lys Ala Thr Leu Gly Leu Gly Gly Val Ala Val Val Leu Gly Ala Val Met Ala Ala Met Gly Phe Phe Ser Tyr Leu Gly Ile Arg Ser Ser Leu Val Ile Leu Gln Val Val Pro Phe Leu Val Leu Ser Val Gly Ala Asp Asn Ile Phe Ile Phe Val Leu Glu Tyr Gln Arg Leu Pro Arg Pro Gly Glu Pro Arg Glu Val His Ile Gly Arg Ala Leu Gly Arg Val Ala Pro Ser Met Leu Leu Cys Ser Leu Ser Glu Ala Ile Cys Phe Phe Leu Gly Ala Leu Thr Pro Met Pro Ala Val Arg Thr Phe Ala Leu Thr Ser Gly Leu Ala Val Ile Leu Asp Phe Leu Leu Gln Met Ser Ala Phe Val Ala Leu Leu Ser Leu Asp Ser Lys Arg Gln Glu Ala Ser Arg Leu Asp Val Cys Cys Cys Val Lys Pro Gln

Glu Leu Pro Pro Gly Gln Gly Glu Gly Leu Leu Gly Phe Phe

- Gln Lys Ala Tyr Ala Pro Phe Leu Leu His Trp Ile Thr Arg Gly Val 835 840 845
- Val Leu Leu Phe Leu Ala Leu Phe Gly Val Ser Leu Tyr Ser Met 850 855 860
- Cys His Ile Ser Val Gly Leu Asp Gln Glu Leu Ala Leu Pro Lys Asp 865 870 875 880
- Ser Tyr Leu Leu Asp Tyr Phe Leu Phe Leu Asn Arg Tyr Phe Glu Val 885 890 895
- Gly Ala Pro Val Tyr Phe Val Thr Thr Leu Gly Tyr Asn Phe Ser Ser 900 905 910
- Glu Ala Gly Met Asn Ala Ile Cys Ser Ser Ala Gly Cys Asn Asn Phe 915 920 925
- Ser Phe Thr Gln Lys Ile Gln Tyr Ala Thr Glu Phe Pro Glu Gln Ser 930 935 940
- Tyr Leu Ala Ile Pro Ala Ser Ser Trp Val Asp Asp Phe Ile Asp Trp 945 950 955 960
- Leu Thr Pro Ser Ser Cys Cys Arg Leu Tyr Ile Ser Gly Pro Asn Lys
 965 970 975
- Asp Lys Phe Cys Pro Ser Thr Val Asn Ser Leu Asn Cys Leu Lys Asn 980 985 990
- Cys Met Ser Ile Thr Met Gly Ser Val Arg Pro Ser Val Glu Gln Phe 995 1000 1005
- His Lys Tyr Leu Pro Trp Phe Leu Asn Asp Arg Pro Asn Ile Lys 1010 1015 1020
- Cys Pro Lys Gly Gly Leu Ala Ala Tyr Ser Thr Ser Val Asn Leu 1025 1030 1035
- Thr Ser Asp Gly Gln Val Leu Ala Ser Arg Phe Met Ala Tyr His 1040 1045 1050
- Lys Pro Leu Lys Asn Ser Gln Asp Tyr Thr Glu Ala Leu Arg Ala 1055 1060 1065

Ala Arg Glu Leu Ala Ala Asn Ile Thr Ala Asp Leu Arg Lys Val Pro Gly Thr Asp Pro Ala Phe Glu Val Phe Pro Tyr Thr Ile Thr Asn Val Phe Tyr Glu Gln Tyr Leu Thr Ile Leu Pro Glu Gly Leu Phe Met Leu Ser Leu Cys Leu Val Pro Thr Phe Ala Val Ser Cys Leu Leu Leu Gly Leu Asp Leu Arg Ser Gly Leu Leu Asn Leu Leu Ser Ile Val Met Ile Leu Val Asp Thr Val Gly Phe Met Ala Leu Trp Asp Ile Ser Tyr Asn Ala Val Ser Leu Ile Asn Leu Val Ser Ala Val Gly Met Ser Val Glu Phe Val Ser His Ile Thr Arg Ser Phe Ala Ile Ser Thr Lys Pro Thr Trp Leu Glu Arg Ala Lys Glu Ala Thr Ile Ser Met Gly Ser Ala Val Phe Ala Gly Val Ala Met Thr Asn Leu Pro Gly Ile Leu Val Leu Gly Leu Ala Lys Ala Gln Leu Ile Gln Ile Phe Phe Phe Arg Leu Asn Leu Leu Ile Thr Leu Leu Gly Leu Leu His Gly Leu Val Phe Leu Pro Val Ile Leu Ser Tyr Val Gly Pro Asp Val Asn Pro Ala Leu Ala Leu Glu Gln Lys Arg Ala Glu Glu Ala Val Ala Val Met Val Ala Ser Cys Pro Asn His Pro Ser Arg Val Ser Thr Ala Asp Asn Ile Tyr Val Asn

1295 1300 1305

His Ser Phe Glu Gly Ser Ile Lys Gly Ala Gly Ala Ile Ser Asn 1310 1315 1320

Phe Leu Pro Asn Asn Gly Arg Gln Phe 1325 1330

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<213> Rattus sp.

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<211> 458

<212> DNA

<213> Rattus sp.

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<210> 7

<211> 896

<212> DNA

<213> Rattus sp.

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<210> 8

<211> 3124

<212> DNA

<213> Rattus sp.

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<211> 4484

<212> DNA

<213> Rattus sp.

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<211> 3993

<212> DNA

<213> Rattus sp.

<220>

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tcg gcc cag ggt gag ctc tac aca ccc act cac aaa gct ggc ttc tgc Ser Ala Gln Gly Glu Leu Tyr Thr Pro Thr His Lys Ala Gly Phe Cys 20 25 30	96
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Thr Phe Tyr Glu Glu Cys Gly Lys Asn Pro Glu Leu Ser Gly Gly Leu 35 40 45	
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Thr Ser Leu Ser Asn Ile Ser Cys Leu Ser Asn Thr Pro Ala Arg His 50 55 60	
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Val Thr Gly Asp His Leu Ala Leu Leu Gln Arg Val Cys Pro Arg Leu 65 70 75 80	
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Tyr Asn Gly Pro Asn Asp Thr Tyr Ala Cys Cys Ser Thr Lys Gln Leu 85 90 95	
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Cys	Ser 130	Pro	Asp	Gln	Ser	Leu 135	Phe	Ile	Asn	Val	Thr 140	Arg	Val	Val	Gln	
	gac Asp															480
	cgc Arg	_		_		_	_				_	_			_	528
	cct Pro															576
	tct Ser															624
	999 Gly 210															672
	ggc Gly															720
	ccc Pro															768
	gac Asp															816
	cct Pro							_				_	_			864
	atc Ile 290			Ala	Val		Val	Leu	Leu		Val					912
	cga Arg															960
	gcc Ala															1008
	ggc Gly															1056
	act Thr	_	_	_	_				_			_	_		_	1104
	ctg Leu															1152

3/0	3/3	360

_			_		_	cgg Arg		_	_	_			_				1200
						acc Thr										÷	1248
		-		_		gac Asp		_	_				_			Š	1296
						gac Asp											1344
	_		_		_	caa Gln 455						_	_	_		-	1392
			_	_		tgc Cys		_				_				-	1440
_				_	_	gtc Val		-						_		3	1488
						ctc Leu										1	1536
						aag Lys										1	1584
		_			_	ggc Gly 535		_	_	-	_	_	_		_	3	1632
						ttc Phe										1	1680
						gca Ala										1	1728
				_	_	gat Asp		_	_	_		_	_			1	1776
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	tac Tyr								_				_	_		2688
	cct Pro															2736
	gca Ala		_		_		_		_	_		_		_		2784
	cta Leu 930															2832
	gtg Val													_		2880
	acc Thr															2928
	gat Asp						-	_					_			2976
	tgc Cys							Pro					r T		aa cag lu Gln	3024
Asn		Met 995 aag Lys	Asn g tao	Arg		Leu tgg	Gly 1000 g tt p Pl	Pro	Val	Arg	g Pro at ac sp Tl	o Thi	r T 05	hr G aac	lu Gln	3024
Asn ttt Phe aga	Cys cat His	Met 995 aag Lys	Asn tac Tyr	Arg c ctg Lei	Thr g ccc	teu tgg Trp 101	Gly 1000 g tt p Pl L5	Pro C ct ne Le	ya] gaa eu As	at gat ag	at acsp Tl	o Thi	r T	hr G aac Asn gtg	lu Gln atc Ile aat	
Asn ttt Phe aga Arg	cat His 1010 tgt Cys 1025	Met 995 aac Lys cct Pro	Asn g tac g tac taaa c Lys	Arg	Thr g ccc g ggc g Gly	Leu tgg Trr 101 ccta Leu 103	Gly 1000 g tt p Ph L5 a go a Al 30	Pro Ca ct Ca go Ca go La Al	o Val	Argat gasan As	at ac sp Tl ga ac rg Tl ag tt	This 100 cg () 100 cg	r T	aac Asn gtg Val	lu Gln atc Ile aat Asn	3069
Asn ttt Phe aga Arg ttg Leu	cat His 1010 tgt Cys 1025 agc Ser	Met 995 aag Lys cct Pro	Asn g tac g tac c aaa c Lys a gat c Asp	Arg	Thr g ccc g ggc / Gly c cac	teu tgg Trr 101 cta Leu 103 gatt 116 cta	Gly 1000 g tt p Ph L5 a go at at E II	Product of the Leader of the L	o Val	at gasn As at ag	at actsp Tl ga acrg Tl ag tt ln pl lca ga	D Th: 100 cg (hr : 020 cc : hr : 035 cc : he : 1050	r T	aac Asn gtg Val gcc Ala	lu Gln atc Ile aat Asn tac Tyr	3069
Asn ttt Phe aga Arg ttg Leu cac His	cat His 1010 tgt Cys 1025 agc Ser 1040 aag	Met 995 aag Lys cct Pro ccc Pro ccc Arg	Asn tac	Arg c ctg c Let a ggg c Gly c gg c Arg	Thr g ccc g ggc / Gly c cac / Glr g aac g Asr	Leu tgg Trg 101 c cta 103 g att 116 c tca 106 a gcc	Gly 1000 H ttp Ph L5 A go A at E Gl 500 C aa A As	Product of the Leader of the L	eg aaeu Aseg ta Ty	at gasn Asat agar Andre Control Contro	at ac sp Tl ga ac rg Tl l l l l l l c a ga r f l l c a ga r f l c a ga r f l c a ga r f l l a ga r f l a ga r	o This 100 cg chr in 100 cc ch	r T	aac Asn gtg Val gcc Ala ctc Leu	lu Gln atc Ile aat Asn tac Tyr cgg Arg	3069 3114 3159
Asn ttt Phe aga Arg ttg Leu cac His gca Ala gtg Val	cat His 1010 tgt Cys 1025 agc Ser 1040 aag Lys 1055	Met 995 aag Lys cct Pro cca Ser cca Pro cca Gly ggs Gly	Asn tags Tyr a a a a a a a a a a a a a a a a a a a	Arg c ctg c Let a ggg c Gl c Ggg c Arg c Let a gat c Asp	Thr g ccc i Pro g ggo / Gly c cac / Glr g aac g Asr a gca i Ala c ccc p Pro	teu tgg Trr 101 c cta 103 d att 104 c tca 106 d Asr 109	Gly 1000 H ttp Ph L5 A go A s Global A	Property of the Leader All and the All and the Salar All and the S	o Val	at gasan Asat agar Asat agar Gi	at ac sp Tl ga ac rg Tl la gt ti la ga rg Tl la gt ti la ga rg Tl	Th: 100 cg chr : 020 cc chr : 035 cc che l 050 ca che l 065 ca che l 080 cc chr : 080 cc chr : 080 cc chr : 080 cc chr : 095	r T	hr G aac Asn gtg Val gcc Ala ctc Leu cgg Arg acg Thr	lu Gln atc Ile aat Asn tac Tyr cgg Arg aag Lys atc Ile	3069 3114 3159 3204

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						aat Asn 1165									3519
						gtg Val 1180							acc Thr		3564
						aag Lys 1195									3609
						ggc Gly 1210									3654
						atc Ile 1225									3699.
	ctt Leu 1235					ttc Phe 1240									3744
		Gly	Leu	Leu	His	ggc Gly 1255	Leu		Phe	Leu		Val			3789
						gtt Val 1270									3834
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Thr Ser Leu Ser Asn Ile Ser Cys Leu Ser Asn Thr Pro Ala Arg His 50 55 60

Val Thr Gly Asp His Leu Ala Leu Leu Gln Arg Val Cys Pro Arg Leu 65 70 75 80

Tyr Asn Gly Pro Asn Asp Thr Tyr Ala Cys Cys Ser Thr Lys Gln Leu 85 90 95

Val Ser Leu Asp Ser Ser Leu Ser Ile Thr Lys Ala Leu Leu Thr Arg 100 105 110

Cys Pro Ala Cys Ser Glu Asn Phe Val Ser Ile His Cys His Asn Thr 115 120 125

Cys Ser Pro Asp Gln Ser Leu Phe Ile Asn Val Thr Arg Val Val Gln 130 135 140

Arg Asp Pro Gly Gln Leu Pro Ala Val Val Ala Tyr Glu Ala Phe Tyr 145 150 155 160

Gln Arg Ser Phe Ala Glu Lys Ala Tyr Glu Ser Cys Ser Arg Val Arg 165 170 175

Ile Pro Ala Ala Ala Ser Leu Ala Val Gly Ser Met Cys Gly Val Tyr 180 185 190 Gly Ser Ala Leu Cys Asn Ala Gln Arg Trp Leu Asn Phe Gln Gly Asp Thr Gly Asn Gly Leu Ala Pro Leu Asp Ile Thr Phe His Leu Leu Glu Pro Gly Gln Ala Leu Ala Asp Gly Met Lys Pro Leu Asp Gly Lys Ile Thr Pro Cys Asn Glu Ser Gln Gly Glu Asp Ser Ala Ala Cys Ser Cys Gln Asp Cys Ala Ala Ser Cys Pro Val Ile Pro Pro Pro Pro Ala Leu Arg Pro Ser Phe Tyr Met Gly Arg Met Pro Gly Trp Leu Ala Leu Ile Ile Ile Phe Thr Ala Val Phe Val Leu Leu Ser Val Val Leu Val Tyr Leu Arg Val Ala Ser Asn Arg Asn Lys Asn Lys Thr Ala Gly Ser Gln Glu Ala Pro Asn Leu Pro Arg Lys Arg Arg Phe Ser Pro His Thr Val Leu Gly Arg Phe Phe Glu Ser Trp Gly Thr Arg Val Ala Ser Trp Pro Leu Thr Val Leu Ala Leu Ser Phe Ile Val Val Ile Ala Leu Ser Val Gly Leu Thr Phe Ile Glu Leu Thr Thr Asp Pro Val Glu Leu Trp Ser Ala Pro Lys Ser Gln Ala Arg Lys Glu Lys Ala Phe His Asp Glu His Phe Gly Pro Phe Phe Arg Thr Asn Gln Ile Phe Val Thr Ala Lys Asn Arg Ser Ser Tyr Lys Tyr Asp Ser Leu Leu Gly Pro Lys Asn Phe

Ser Gly Ile Leu Ser Leu Asp Leu Leu Gln Glu Leu Glu Leu Gln Glu Arg Leu Arg His Leu Gln Val Trp Ser His Glu Ala Gln Arg Asn Ile Ser Leu Gln Asp Ile Cys Tyr Ala Pro Leu Asn Pro His Asn Thr Ser Leu Thr Asp Cys Cys Val Asn Ser Leu Leu Gln Tyr Phe Gln Asn Asn His Thr Leu Leu Leu Thr Ala Asn Gln Thr Leu Asn Gly Gln Thr Ser Leu Val Asp Trp Lys Asp His Phe Leu Tyr Cys Ala Asn Ala Pro Leu Thr Tyr Lys Asp Gly Thr Ala Leu Ala Leu Ser Cys Ile Ala Asp Tyr Gly Ala Pro Val Phe Pro Phe Leu Ala Val Gly Gly Tyr Gln Gly Thr Asp Tyr Ser Glu Ala Glu Ala Leu Ile Ile Thr Phe Ser Ile Asn Asn Tyr Pro Ala Asp Asp Pro Arg Met Ala His Ala Lys Leu Trp Glu Glu Ala Phe Leu Lys Glu Met Gln Ser Phe Gln Arg Ser Thr Ala Asp Lys Phe Gln Ile Ala Phe Ser Ala Glu Arg Ser Leu Glu Asp Glu Ile Asn Arg Thr Thr Ile Gln Asp Leu Pro Val Phe Ala Ile Ser Tyr Leu Ile Val Phe Leu Tyr Ile Ser Leu Ala Leu Gly Ser Tyr Ser Arg Trp Ser Arg Val Ala Val Asp Ser Lys Ala Thr Leu Gly Leu Gly Gly

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Glu Ala Gly Met Asn Ala Ile Cys Ser Ser Ala Gly Cys Glu Ser Phe

Ser Leu Thr Gln Lys Ile Gln Tyr Ala Ser Glu Phe Pro Asn Gln Ser

- Tyr Val Ala Ile Ala Ala Ser Ser Trp Val Asp Asp Phe Ile Asp Trp
- Leu Thr Pro Ser Ser Ser Cys Cys Arg Ile Tyr Thr Arg Gly Pro His
- Lys Asp Glu Phe Cys Pro Ser Thr Asp Thr Ser Phe Asn Cys Leu Lys
- Asn Cys Met Asn Arg Thr Leu Gly Pro Val Arg Pro Thr Thr Glu Gln
- Phe His Lys Tyr Leu Pro Trp Phe Leu Asn Asp Thr Pro Asn Ile
- Arg Cys Pro Lys Gly Gly Leu Ala Ala Tyr Arg Thr Ser Val Asn
- Leu Ser Ser Asp Gly Gln Ile Ile Ala Ser Gln Phe Met Ala Tyr
- His Lys Pro Leu Arg Asn Ser Gln Asp Phe Thr Glu Ala Leu Arg
- Ala Ser Arg Leu Leu Ala Ala Asn Ile Thr Ala Glu Leu Arg Lys
- Val Pro Gly Thr Asp Pro Asn Phe Glu Val Phe Pro Tyr Thr Ile
- Ser Asn Val Phe Tyr Gln Gln Tyr Leu Thr Val Leu Pro Glu Gly
- Ile Phe Thr Leu Ala Leu Cys Phe Val Pro Thr Phe Val Val Cys
- Tyr Leu Leu Gly Leu Asp Ile Arg Ser Gly Ile Leu Asn Leu
- Leu Ser Ile Ile Met Ile Leu Val Asp Thr Ile Gly Leu Met Ala

Val Trp Gly Ile Ser Tyr Asn Ala Val Ser Leu Ile Asn Leu Val 1160 1165 1170

Thr Ala Val Gly Met Ser Val Glu Phe Val Ser His Ile Thr Arg 1175 1180 1185

Ser Phe Ala Val Ser Thr Lys Pro Thr Arg Leu Glu Arg Ala Lys 1190 1195 1200

Asp Ala Thr Ile Phe Met Gly Ser Ala Val Phe Ala Gly Val Ala 1205 1210 1215

Met Thr Asn Phe Pro Gly Ile Leu Ile Leu Gly Phe Ala Gln Ala 1220 1225 1230

Gln Leu Ile Gln Ile Phe Phe Phe Arg Leu Asn Leu Leu Ile Thr 1235 1240 1245

Leu Leu Gly Leu Leu His Gly Leu Val Phe Leu Pro Val Val Leu 1250 1260

Ser Tyr Leu Gly Pro Asp Val Asn Gln Ala Leu Val Leu Glu Glu 1265 1270 1275

Lys Leu Ala Thr Glu Ala Ala Met Val Ser Glu Pro Ser Cys Pro 1280 1285 1290

Gln Tyr Pro Phe Pro Ala Asp Ala Asn Thr Ser Asp Tyr Val Asn 1295 1300 1305

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110

105

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							ggc Gly							635
_		_		_	-	_	tgg Trp				_	 _		683
							atc Ile							731
				_			cag Gln		_			 _	_	779
							gac Asp 250							827
							ata Ile							875
							ccg Pro							923
							gtc Val							971
							agc Ser							1019
				_	_		agc Ser 330							1067
							acg Thr							1115

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				999 Gly 710								_				2219
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				ttc Phe												2363
				ctg Leu												2411
				gcc Ala 790												2459
				cgg Arg												2507
				ggc Gly												2555
				ccc Pro												2603

835 840 845

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cac atc agc gtg gga ctg gac cag gag ctg gcc ctg ccc aag gac tcg His Ile Ser Val Gly Leu Asp Gln Glu Leu Ala Leu Pro Lys Asp Ser 870 875 880	2699
tac ctg ctt gac tat ttc ctc ttt ctg aac cgc tac ttc gag gtg ggg Tyr Leu Leu Asp Tyr Phe Leu Phe Leu Asn Arg Tyr Phe Glu Val Gly 885 890 895	2747
gcc ccg gtg tac ttt gtt acc acc ttg ggc tac aac ttc tcc agc gag Ala Pro Val Tyr Phe Val Thr Thr Leu Gly Tyr Asn Phe Ser Ser Glu 900 905 910	2795
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ccg Pro 1115	Ala				ttc Phe 1120										3443
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	Cys				acc Thr 1150										3533
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	Ile				ggc Gly 1255										3848
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cat His 1280	Gly				ctg Leu 1285										3938
	Val				ctg Leu 1300										3983

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Ala Arg Cys Asn Glu Ser Gln Gly Asp Asp Val Ala Thr Cys Ser Cys

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- Ile Ile Leu Cys Ser Val Phe Ala Val Val Thr Ile Leu Leu Val Gly 290 295 300
- Phe Arg Val Ala Pro Ala Arg Asp Lys Ser Lys Met Val Asp Pro Lys 305 310 315 320
- Lys Gly Thr Ser Leu Ser Asp Lys Leu Ser Phe Ser Thr His Thr Leu 325 330 335
- Leu Gly Gln Phe Phe Gln Gly Trp Gly Thr Trp Val Ala Ser Trp Pro 340 345 350
- Leu Thr Ile Leu Val Leu Ser Val Ile Pro Val Val Ala Leu Ala Ala 355 360 365
- Gly Leu Val Phe Thr Glu Leu Thr Thr Asp Pro Val Glu Leu Trp Ser 370 380
- Ala Pro Asn Ser Gln Ala Arg Ser Glu Lys Ala Phe His Asp Gln His 385 390 395 400
- Phe Gly Pro Phe Phe Arg Thr Asn Gln Val Ile Leu Thr Ala Pro Asn 405 410 415
- Arg Ser Ser Tyr Arg Tyr Asp Ser Leu Leu Leu Gly Pro Lys Asn Phe 420 425 430
- Ser Gly Ile Leu Asp Leu Asp Leu Leu Leu Glu Leu Glu Leu Gln 435 440 445
- Glu Arg Leu Arg His Leu Gln Val Trp Ser Pro Glu Ala Gln Arg Asn 450 455 460
- Ile Ser Leu Gln Asp Ile Cys Tyr Ala Pro Leu Asn Pro Asp Asn Thr 465 470 475 480
- Ser Leu Tyr Asp Cys Cys Ile Asn Ser Leu Leu Gln Tyr Phe Gln Asn 495

Asn Arg Thr Leu Leu Leu Thr Ala Asn Gln Thr Leu Met Gly Gln Thr Ser Gln Val Asp Trp Lys Asp His Phe Leu Tyr Cys Ala Asn Ala Pro Leu Thr Phe Lys Asp Gly Thr Ala Leu Ala Leu Ser Cys Met Ala Asp Tyr Gly Ala Pro Val Phe Pro Phe Leu Ala Ile Gly Gly Tyr Lys Gly Lys Asp Tyr Ser Glu Ala Glu Ala Leu Ile Met Thr Phe Ser Leu Asn Asn Tyr Pro Ala Gly Asp Pro Arg Leu Ala Gln Ala Lys Leu Trp Glu Glu Ala Phe Leu Glu Glu Met Arg Ala Phe Gln Arg Arg Met Ala Gly Met Phe Gln Val Thr Phe Met Ala Glu Arq Ser Leu Glu Asp Glu Ile Asn Arg Thr Thr Ala Glu Asp Leu Pro Ile Phe Ala Thr Ser Tyr Ile Val Ile Phe Leu Tyr Ile Ser Leu Ala Leu Gly Ser Tyr Ser Ser Trp Ser Arg Val Met Val Asp Ser Lys Ala Thr Leu Gly Leu Gly Gly Val Ala Val Val Leu Gly Ala Val Met Ala Ala Met Gly Phe Phe Ser Tyr Leu Gly Ile Arg Ser Ser Leu Val Ile Leu Gln Val Val Pro Phe Leu Val Leu Ser Val Gly Ala Asp Asn Ile Phe Ile Phe Val Leu Glu Tyr Gln Arg Leu Pro Arg Arg Pro Gly Glu Pro Arg Glu Val His Ile

Gly Arg Ala Leu Gly Arg Val Ala Pro Ser Met Leu Leu Cys Ser Leu Ser Glu Ala Ile Cys Phe Phe Leu Gly Ala Leu Thr Pro Met Pro Ala Val Arg Thr Phe Ala Leu Thr Ser Gly Leu Ala Val Ile Leu Asp Phe Leu Leu Gln Met Ser Ala Phe Val Ala Leu Leu Ser Leu Asp Ser Lys Arg Gln Glu Ala Ser Arg Leu Asp Val Cys Cys Cys Val Lys Pro Gln Glu Leu Pro Pro Gly Gln Gly Glu Gly Leu Leu Gly Phe Phe Gln Lys Ala Tyr Ala Pro Phe Leu Leu His Trp Ile Thr Arg Gly Val Val Leu Leu Phe Leu Ala Leu Phe Gly Val Ser Leu Tyr Ser Met Cys His Ile Ser Val Gly Leu Asp Gln Glu Leu Ala Leu Pro Lys Asp Ser Tyr Leu Leu Asp Tyr Phe Leu Phe Leu Asn Arg Tyr Phe Glu Val Gly Ala Pro Val Tyr Phe Val Thr Thr Leu Gly Tyr Asn Phe Ser Ser Glu Ala Gly Met Asn Ala Ile Cys Ser Ser Ala Gly Cys Asn Asn Phe Ser Phe Thr Gln Lys Ile Gln Tyr Ala Thr Glu Phe Pro Glu Gln Ser Tyr Leu Ala Ile Pro Ala Ser Ser Trp Val Asp Asp Phe Ile Asp Trp Leu Thr Pro Ser Ser Cys Cys Arg Leu Tyr Ile Ser Gly Pro Asn Lys

- Asp Lys Phe Cys Pro Ser Thr Val Asn Ser Leu Asn Cys Leu Lys Asn 980 985 990
- Cys Met Ser Ile Thr Met Gly Ser Val Arg Pro Ser Val Glu Gln Phe 995 1000 1005
- His Lys Tyr Leu Pro Trp Phe Leu Asn Asp Arg Pro Asn Ile Lys 1010 1015 1020
- Cys Pro Lys Gly Gly Leu Ala Ala Tyr Ser Thr Ser Val Asn Leu 1025 1030 1035
- Thr Ser Asp Gly Gln Val Leu Asp Thr Val Ala Ile Leu Ser Pro 1040 1045 1050
- Arg Leu Glu Tyr Ser Gly Thr Ile Ser Ala His Cys Asn Leu Tyr 1055 1060 1065
- Leu Leu Asp Ser Ala Ser Arg Phe Met Ala Tyr His Lys Pro Leu 1070 1080
- Lys Asn Ser Gln Asp Tyr Thr Glu Ala Leu Arg Ala Ala Arg Glu 1085 1090 1095
- Leu Ala Ala Asn Ile Thr Ala Asp Leu Arg Lys Val Pro Gly Thr 1100 1105 1110
- Asp Pro Ala Phe Glu Val Phe Pro Tyr Thr Ile Thr Asn Val Phe 1115 1120 1125
- Tyr Glu Gln Tyr Leu Thr Ile Leu Pro Glu Gly Leu Phe Met Leu 1130 1135 1140
- Ser Leu Cys Leu Val Pro Thr Phe Ala Val Ser Cys Leu Leu Leu 1145 1150 1155
- Gly Leu Asp Leu Arg Ser Gly Leu Leu Asn Leu Leu Ser Ile Val 1160 1165 1170
- Met Ile Leu Val Asp Thr Val Gly Phe Met Ala Leu Trp Gly Ile 1175 1180 1185
- Ser Tyr Asn Ala Val Ser Leu Ile Asn Leu Val Ser Ala Val Gly 1190 1195 1200
- Met Ser Val Glu Phe Val Ser His Ile Thr Arg Ser Phe Ala Ile

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Pro	Gly 1250	Ile	Leu	Val	Leu	Gly 1255	Leu	Ala	Lys	Ala	Gln 1260	Leu	Ile	Gln
Ile	Phe 1265	Phe	Phe	Arg	Leu	Asn 1270	Leu	Leu	Ile	Thr	Leu 1275	Leu	Gly	Leu
Leu	His 1280	Gly	Leu	Val	Phe	Leu 1285	Pro	Val	Ile	Leu	Ser 1290	Tyr	Val	Gly
Pro	Asp 1295	Val	Asn	Pro	Ala	Leu 1300	Ala	Leu	Glu	Gln	Lys 1305	Arg	Ala	Glu
Glu	Ala 1310	Val	Ala	Ala	Val	Met 1315	Val	Ala	Ser	Cys	Pro 1320	Asn	His	Pro
Ser	Arg 1325	Val	Ser	Thr	Ala	Asp 1330	Asn	Ile	Tyr	Val	Asn 1335	His	Ser	Phe
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